Appl. No. Filed

09/853,080

May 9, 2001

()2

possibly deleted from one or more amino acids from the heptapeptide Glu-Thr-Lys-Ser-Trp-Phe-Thr or from the tripeptide Cys-Arg-Ala,

Please insert the following paragraph after the paragraph on page 9, line 23 ("SEQ ID NO:12:):

 $0^{2}$ 

Glu Gly Ser Leu Ala Lys Glu Lys Thr Gln Thr Leu,

Please replace the paragraph on page 10, lines 6-7, with the following rewritten paragraph:

4

Asp Ser Cys Pro Glu Glu Pro Gln Leu Arg Met Lys Asn Asn Glu Glu Ala Glu Asp Tyr Asp Asp Asp Leu Thr Asp Ser Glu Met,

Please replace the paragraph on page 10, lines 18-20, with the following rewritten paragraph:

OS.

Asp Asp Arg Ser Tyr Lys Ser Gln Tyr Leu Asn Asn Gly Pro Gln Arg Ile Gly Arg Lys Tyr Lys Lys,

Please replace the paragraph on page 11, line 8, with the following rewritten paragraph:

do

Ser Cys Asp Lys Asn Thr Gly Asp Tyr Tyr Gly Asp Ser Tyr Glu Asp,

Please replace the paragraph on page 21, line 12, with the following rewritten paragraph:

Ser Cys Asp Lys Asn Thr Gly Asp Tyr Tyr Gly Asp Ser Tyr Glu Asp,

Please replace the paragraph beginning at page 36, line 2, with the following rewritten paragraph:

8

In the clotting test, significant inhibition of FVIII activity was recorded in the presence of rabbit anti-(Cys<sup>329</sup>-Asp<sup>348</sup>) and anti-(Arg<sup>1652</sup>-Tyr<sup>1664</sup>) antibodies, but different inhibition patterns were observed. Inhibition by anti-(Arg<sup>1652</sup>-Tyr<sup>1664</sup>) follows second-order kinetics with a drastic reduction in FVIII activity. Anti-(Cys<sup>329</sup>-Asp<sup>348</sup>) Ig is less efficient and shows a more complex type of reaction, with a non-linear dependence on the antibody concentration. Epitope Arg<sup>1652</sup>-Tyr<sup>1664</sup> and the adjacent major binding site vWF (residues Glu<sup>1675</sup>-Glu<sup>1684</sup>) are located in the acidic